

Application Ser. No.: 10/613,588

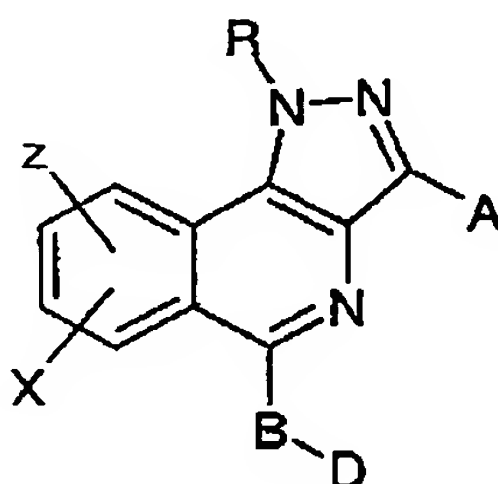
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Examiner: Huang, Evelyn Mei

**Amendment Pursuant to 37 C.F.R. § 1.121****IN THE CLAIMS:**

The claims set forth below with amendments as indicated will replace all prior versions and listing of claims in the application.

1. (currently amended) A compound of the formula I



(I)

or a stereoisomeric form or a pharmaceutically acceptable salt of the compound of the formula I, wherein

A is  $-(C_1-C_6)$ -alkyl, in which alkyl is straight-chain or branched and is optionally substituted, once or more, independently of each other, by

$-O-R^1$  or

$-C(O)-OR^1$ ,

$-C(O)-NR^1R^1$ ,

$-C(O)-NR^1-SO_2R^1$ ,

$-NR^1R^1$ ,

$-CN$ , in which  $R^1$  is

hydrogen,

$-(C_1-C_6)$ -alkyl,

$-(C_6-C_{14})$ aryl or

fluoroalkyl of the formula  $-C_nH_xF_y$  or fluoroalkoxy of the formula  $-OC_nH_xF_y$ , wherein  $n$  is an integer from 1 to 6,  $x$  is an integer from 0 to 12,  $y$  is an integer from 1 to 13 and sum of  $x$  and  $y$  is  $2n + 1$ ,

$-O-R^1$ ,

$-SR^1$ ,

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-S(O)-R<sup>1</sup>

-S(O)<sub>2</sub>-R<sup>1</sup>

-C(O)-OR<sup>1</sup>,

fluoroalkyl of the formula -C<sub>n</sub>H<sub>x</sub>F<sub>y</sub> or fluoroalkoxy of the formula

-OC<sub>n</sub>H<sub>x</sub>F<sub>y</sub>, wherein n is an integer from 1 to 6, x is an integer from 0 to

12, y is an integer from 1 to 13 and sum of x and y is 2n + 1,

-C(O)-NR<sup>1</sup>R<sup>1</sup>,

-C(O)-NR<sup>1</sup>-SO<sub>2</sub>R<sup>1</sup>,

-NR<sup>1</sup>R<sup>1</sup>,

-CN,

monocyclic or polycyclic heteroaryl having from 5 to 14 ring members, which contains 1 to 5 heteroatoms as ring members, wherein

heteroatoms are selected from N, O and S, and in which heteroaryl is unsubstituted or optionally substituted once or more, independently of each other, by R<sup>2</sup>, in which R<sup>2</sup> is

-(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

-OH,

-O-(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

halogen,

-N(R<sup>3</sup>)-R<sup>4</sup>, in which R<sup>3</sup> and R<sup>4</sup> are, independently of each other, hydrogen atom or -(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

fluoroalkyl of the formula -C<sub>n</sub>H<sub>x</sub>F<sub>y</sub> or fluoroalkoxy of the formula -OC<sub>n</sub>H<sub>x</sub>F<sub>y</sub>, wherein n is an integer from 1 to 4, x is an integer from 0 to 8, y is an integer from 1 to 9 and sum of x and y is 2n + 1,

-CN,

-SR<sup>1</sup>,

-S(O)-R<sup>1</sup>,

-S(O)<sub>2</sub>-R<sup>1</sup> or

-C(O)-NR<sup>1</sup>R<sup>1</sup>,

-(C<sub>3</sub>-C<sub>6</sub>)-cycloalkyl, in which cycloalkyl is unsubstituted or substituted, once or more, independently of each other, by R<sup>2</sup>, and R<sup>2</sup> is defined as above,

monocyclic or bicyclic heterocycle having from 5 to 12 ring members, which contains 1 to 5 heteroatoms as ring members, wherein

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heteroatoms are selected from N, O and S, and in which heterocycle is unsubstituted or substituted, once or more, independently of each other, by R<sup>2</sup>, and R<sup>2</sup> is defined as above,

B is a covalent bond,  
-C=CR<sup>1</sup>-,  
—C≡C—,  
-O(CH<sub>2</sub>)<sub>a</sub>-, in which a is an integer from 1 to 4,  
O, S, NR<sup>2</sup>-, -C(O)-, -NR<sup>2</sup>-C(O)-, -C(O)-NR<sup>2</sup>-, -NR<sup>2</sup>-SO<sub>2</sub>-,  
-SO<sub>2</sub>-NR<sup>2</sup>-, -NR<sup>2</sup>-C(O)-NR<sup>2</sup>-, and R<sup>2</sup> is defined as above, or  
-(C<sub>1</sub>-C<sub>4</sub>)-alkylene, in which alkylene is straight-chain or branched and is optionally substituted, once or more, independently of each other, by R<sup>1</sup>, and R<sup>1</sup> is defined as above,

D is -(C<sub>1</sub>-C<sub>6</sub>)-alkyl, in which alkyl is straight-chain or branched and is optionally substituted, once or more, independently of each other, by R<sup>1</sup>, and R<sup>1</sup> is defined as above,

monocyclic or polycyclic heteroaryl having from 5 to 14 ring members, which contains 1 to 5 heteroatoms as ring members, and in which heteroaryl is unsubstituted or is substituted once or more, independently of each other, by R<sup>2</sup>, and R<sup>2</sup> is defined as above,

monocyclic or bicyclic heterocycle having from 5 to 12 ring members, which contains 1 to 5 heteroatoms as ring members, and in which heterocycle is unsubstituted or substituted, once or more, independently of each other, by R<sup>2</sup>, and R<sup>2</sup> is defined as above,

-(C<sub>6</sub>-C<sub>14</sub>)-aryl, in which aryl is unsubstituted or substituted, once or more, independently of each other, by R<sup>2</sup>, and R<sup>2</sup> is defined as above, or

-(C<sub>3</sub>-C<sub>6</sub>)-cycloalkyl, in which cycloalkyl is unsubstituted or substituted, once or more, independently of each other, by R<sup>2</sup>, and R<sup>2</sup> is defined as above, or

B-D is hydrogen,  
halogen,

fluoroalkyl of the formula -C<sub>n</sub>H<sub>x</sub>F<sub>y</sub> or fluoroalkoxy of the formula -OC<sub>n</sub>H<sub>x</sub>F<sub>y</sub>, wherein n is an integer from 1 to 4, x is

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an integer from 0 to 8, y is an integer from 1 to 9 and sum of x and y is  $2n + 1$ ,

$-(CH_2)_a-Y-R^3$ , in which a is an integer from 1 to 4, Y is O, S,  $NR^2$ , and  $R^3$  is

$-(C_1-C_6)$ -alkyl,  
 $-(C_6-C_{14})$ -aryl,  
 $-(C_3-C_6)$ -cycloalkyl, and

R is hydrogen,  
 $-(C_1-C_6)$ -alkyl, or  
 $-(C_6-C_{14})$ -aryl- $-(C_1-C_6)$ -alkyl, in which aryl is unsubstituted or substituted, once or more, independently of each other, by  $R^2$ , and  $R^2$  is defined as above, and

X and Z are identical or different and are, independently of each other selected from:

hydrogen atom,  
 $-(C_1-C_4)$ -alkyl,  
-OH,  
-O- $-(C_1-C_4)$ -alkyl),  
halogen,  
fluoroalkyl of the formula  $-C_nH_xF_y$  or fluoroalkoxy of the formula  $-OC_nH_xF_y$ , wherein n is an integer from 1 to 6, x is an integer from 0 to 12, y is an integer from 1 to 13 and sum of x and y is  $2n + 1$ ,  
 $-C(O)-OR^1$ ,  
 $-C(O)-NR^1R^1$ ,  
 $-C(O)-NR^1-SO_2R^1$ ,  
 $-NR^1R^1$ ,  
 $-NR^1-C(O)-NR^1R^1$ ,  
 $-NR^1-C(O)-R^1$ ,  
 $-NR^1-C(O)-OR^1$ ,  
 $-O-C(O)-NR^1R^1$ ,  
-CN,  
-SR<sup>1</sup>,  
-S(O)-R<sup>1</sup>,  
-S(O)<sub>2</sub>-R<sup>1</sup>,  
-S(O)<sub>2</sub>-NR<sup>1</sup>R<sup>1</sup>,  
-NR<sup>1</sup>-SO<sub>2</sub>-R<sup>1</sup>, in which R<sup>1</sup> is as defined above,

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monocyclic or bicyclic heterocycle having from 5 to 12 ring members, which contains 1 to 5 heteroatoms as ring members, wherein heteroatoms are selected from N, O and S, and in which heterocycle is unsubstituted or substituted, once or more, independently of each other, by  $R^2$ , and  $R^2$  is defined as above, or

-(C<sub>3</sub>-C<sub>6</sub>)-cycloalkyl, in which cycloalkyl is unsubstituted or substituted, once or more, independently of each other, by  $R^2$ , and  $R^2$  is defined as above,

with the proviso that when A is -(C<sub>1</sub>-C<sub>6</sub>)-alkyl, in which alkyl is straight-chain or branched and is optionally substituted, once or more, independently of each other, by

-O-R<sup>1</sup> or

-C(O)-OR<sup>1</sup>, in which R<sup>1</sup> is

hydrogen,

-(C<sub>1</sub>-C<sub>6</sub>)-alkyl,

-O-R<sup>1</sup>, in which R<sup>1</sup> is as defined in this proviso,

-C(O)-OR<sup>1</sup>, in which R<sup>1</sup> is as defined in this proviso, or

monocyclic or polycyclic heteroaryl having from 5 to 14 ring members, which contains 1 to 5 heteroatoms as ring members, wherein heteroatoms are selected from N, O and S, and in which heteroaryl is unsubstituted or optionally substituted once or more, independently of each other, by  $R^2$ , in which  $R^2$  is

hydrogen,

-(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

-OH,

-O-(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

halogen, or

-N(R<sup>3</sup>)-R<sup>4</sup>, in which R<sup>3</sup> and R<sup>4</sup> are, independently of each other, hydrogen atom or -(C<sub>1</sub>-C<sub>4</sub>)-alkyl, then at least one of the following applies:

B is not a covalent bond or -(C<sub>1</sub>-C<sub>4</sub>)-alkylene,

~~D is not heteroaryl, heterocycle, -(C<sub>6</sub>-C<sub>14</sub>)-aryl, -(C<sub>3</sub>-C<sub>6</sub>)-cycloalkyl, or~~

~~X and Z are not -(C<sub>1</sub>-C<sub>4</sub>)-alkyl, -OH, -O-(C<sub>1</sub>-C<sub>4</sub>)-alkyl, or halogen.~~

2. (original) A compound of the formula I as claimed in claim 1, wherein

A is -(C<sub>1</sub>-C<sub>3</sub>)-alkyl, in which alkyl is straight-chain or branched and is optionally substituted, once or more, independently of each other, by

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-O-R<sup>1</sup>, or  
 -C(O)-OR<sup>1</sup>, in which R<sup>1</sup> is  
 hydrogen,  
 -(C<sub>1</sub>-C<sub>3</sub>)-alkyl, or  
 -CF<sub>3</sub>

fluoroalkyl of the formula -C<sub>n</sub>H<sub>x</sub>F<sub>y</sub> or fluoroalkoxy of the formula  
 -OC<sub>n</sub>H<sub>x</sub>F<sub>y</sub>, wherein n is an integer from 1 to 3, x is an integer from 0  
 to 6, y is an integer from 1 to 7 and sum of x and y is 2n + 1,

B is a covalent bond or O,

D is phenyl or naphthyl, in which phenyl or naphthyl is unsubstituted  
 or substituted, once or more, independently of each other, by R<sup>2</sup>, in  
 which R<sup>2</sup> is

fluorine, chlorine or bromine,  
 -OH,  
 -CF<sub>3</sub>,  
 -SR<sup>1</sup>, in which R<sup>1</sup> is defined as above,  
 -(C<sub>1</sub>-C<sub>4</sub>)-alkyl

-O-(C<sub>1</sub>-C<sub>2</sub>)-alkyl or  
 -N(R<sup>3</sup>)-R<sup>4</sup>, in which R<sup>3</sup> and R<sup>4</sup> are, independently of  
 each other, hydrogen atom or -(C<sub>1</sub>-C<sub>3</sub>)-alkyl,  
 heteroaryl selected from the group consisting of  
 pyridyl, furanyl, pyrrolyl, isoxazolyl, benzofuranyl, benzothiophenyl,  
 quinolinyl, isoquinolinyl, quinoxalinyl and thiophenyl, in which  
 heteroaryl is unsubstituted or substituted, once or more,  
 independently of each other, by R<sup>2</sup>, in which R<sup>2</sup> is defined as above  
 or

-(C<sub>4</sub>-C<sub>6</sub>)-cycloalkyl, in which cycloalkyl is  
 unsubstituted or substituted, once or more, independently of each  
 other, by R<sup>2</sup>, and R<sup>2</sup> is defined as above, or

B-D is ((CH<sub>2</sub>)<sub>a</sub>-Y-R<sup>3</sup>, in which a is an integer from 1 to 2, Y is O and R<sup>3</sup>  
 is -(C<sub>1</sub>-C<sub>3</sub>)-alkyl, and

R is hydrogen,

-(C<sub>1</sub>-C<sub>3</sub>)-alkyl, or  
 -phenyl-(C<sub>1</sub>-C<sub>3</sub>)-alkyl, and

X and Z are identical or different and are, independently of each  
 other, hydrogen, -C(O)-O(C<sub>1</sub>-C<sub>3</sub>)alkyl, -OCH<sub>3</sub>, -N(CH<sub>3</sub>)<sub>2</sub> or halogen.

3. (original) A compound of the formula 1 as claimed in claim 1, wherein the  
 compound of the formula 1 is selected from the group consisting of:

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5-pyridin-2-yl-3-trifluoromethyl-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(2-trifluoromethyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(3-trifluoromethyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(4-trifluoromethyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
1,3-dimethyl-5-(3-trifluoromethylphenyl)-1H-pyrazolo[4,3-c]-isoquinoline,  
5-phenyl-3-trifluoromethyl-1H-pyrazolo[4,3-c]isoquinoline,  
1,3-dimethyl-5-(3-trifluoromethylphenyl)-1H-pyrazolo[4,3-c]-isoquinoline,  
1,3-dimethyl-5-(2,6-difluorophenyl)-1H-pyrazolo[4,3-c]-isoquinoline,  
1-benzyl-5-cyclohexyl-3-methyl-1H-pyrazolo[4,3-c]-isoquinoline,  
1-benzyl-5-naphthyl-3-methyl-1H-pyrazolo[4,3-c]-isoquinoline,  
5-methoxymethyl-3-methyl-1H-pyrazolo[4,3-c]-isoquinoline,  
7-methoxycarbonyl-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]-isoquinoline,  
7-methoxycarbonyl-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]-  
isoquinoline,  
7-dimethylamino-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
7-dimethylamino-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]-isoquinoline,  
6-dimethylamino-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
6-dimethylamino-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]-isoquinoline,  
8-dimethylamino-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
8-dimethylamino-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]-isoquinoline,  
1,3-dimethyl-5-(3-methyl-thiophen-2-yl)-1H-pyrazolo[4,3-c]-isoquinoline,  
3-methyl-5-phenyl-9-trifluoromethyl-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-pyridin-2-yl-9-trifluoromethyl-1H-pyrazolo[4,3-c]-isoquinoline,  
and  
3-methyl-5-(2,3,4,5,6-pentafluoro-phenyl)-1H-pyrazolo[4,3-c]-isoquinoline.

4. (currently amended) A compound selected from the group consisting of:

5-benzo[b]thiophene-2yl-3-methyl-1H-pyrazolo[4,3-c]isoquinoline,  
7-bromo-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
7-bromo-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,



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5-(3-chloro-benzo[b]thiophen-2-yl)-3-methyl-1H-pyrazolo[4,3-c]-isoquinoline,  
5-(2-chloro-pyridin-3-yl)-3-methyl-1H-pyrazolo[4,3-c]isoquinoline,  
5-(6-chloro-pyridin-3-yl)-3-methyl-1H-pyrazolo[4,3-c]isoquinoline,  
3-ethyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
3-ethyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
3-ethyl-5-pyridin-3-yl-1H-pyrazolo[4,3-c]isoquinoline,  
5-furan-2-yl-3-methyl-1H-pyrazolo[4,3-c]isoquinoline,  
7-methoxy-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(3-methyl-benzofuran-2-yl)-1H-pyrazolo[4,3-c]-isoquinoline,  
~~3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,~~  
6-chloro-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
6-fluoro-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
8-chloro-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
8-fluoro-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
9-chloro-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
9-fluoro-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
3,9-dimethyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(2-methyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(3-methyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
~~3-methyl-5-(4-methyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,~~  
3-methyl-5-(2-bromo-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(3-bromo-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(2-chloro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
~~3-methyl-5-(4-chloro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,~~  
3-methyl-5-(2,4-dichloro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(3,4-dichloro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(2,6-dichloro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(2-fluoro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(4-fluoro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,



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3-methyl-5-(2,4-difluoro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(2,6-difluoro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
6-chloro-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
6-fluoro-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
8-chloro-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
8-fluoro-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
9-chloro-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
9-fluoro-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
3,9-dimethyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(1-methyl-1H-pyrrol-2-yl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-quinolin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-quinolin-3-yl-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-quinoxalin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-thiophen-2-yl-1H-pyrazolo[4,3-c]isoquinoline, and  
3-methyl-5-(3-methyl-thiophen-2-yl)-1H-pyrazolo[4,3-c]isoquinoline.

5. (original) A pharmaceutical composition comprising a therapeutically effective content of at least one compound of the formula I as claimed in claim 1 together with a pharmaceutically suitable carrier optionally in combination with a suitable additive, other active compounds and auxiliary substances.
6. (original) A method of treating a disease condition associated with the increased activity of NIK comprising administering to a patient suffering from said disease condition a therapeutically effective amount of a compound according to claim 1.
7. (original) The method as claimed in claim 6, wherein the compound is according to claim 2.

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8. (original) The method as claimed in claim 6 wherein said compound is selected from the group consisting of:
- 5-pyridin-2-yl-3-trifluoromethyl-1H-pyrazolo[4,3-c]isoquinoline,
  - 3-methyl-5-(2-trifluoromethyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,
  - 3-methyl-5-(3-trifluoromethyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,
  - 3-methyl-5-(4-trifluoromethyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,
  - 1,3-dimethyl-5-(3-trifluoromethylphenyl)-1H-pyrazolo[4,3-c]-isoquinoline,
  - 5-phenyl-3-trifluoromethyl-1H-pyrazolo[4,3-c]isoquinoline,
  - 1,3-dimethyl-5-(3-trifluoromethylphenyl)-1H-pyrazolo[4,3-c]-isoquinoline,
  - 1,3-dimethyl-5-(2,6-difluorophenyl)-1H-pyrazolo[4,3-c]-isoquinoline,
  - 1-benzyl-5-cyclohexyl-3-methyl-1H-pyrazolo[4,3-c]-isoquinoline,
  - 1-benzyl-5-naphthyl-3-methyl-1H-pyrazolo[4,3-c]-isoquinoline,
  - 5-methoxymethyl-3-methyl-1H-pyrazolo[4,3-c]-isoquinoline,
  - 7-methoxycarbonyl-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]-isoquinoline,
  - 7-methoxycarbonyl-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]-isoquinoline,
  - 7-dimethylamino-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,
  - 7-dimethylamino-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]-isoquinoline,
  - 1,3-dimethyl-5-(3-methyl-thiophen-2-yl)-1H-pyrazolo[4,3-c]-isoquinoline,
  - 3-methyl-5-phenyl-9-trifluoromethyl-1H-pyrazolo[4,3-c]isoquinoline,
  - 3-methyl-5-pyridin-2-yl-9-trifluoromethyl-1H-pyrazolo[4,3-c]-isoquinoline,
  - and
  - 3-methyl-5-(2,3,4,5,6-pentafluoro-phenyl)-1H-pyrazolo[4,3-c]-isoquinoline.
9. (original) The method as claimed in claim 6, wherein the disease condition is caused due to an inflammatory component.
10. (original) The method as claimed in claim 6, wherein the diseases are osteoarthritis, rheumatoid arthritis, asthma, irritable bowel disease, Alzheimer's disease, stroke, diabetes, atherosclerosis, multiple sclerosis,

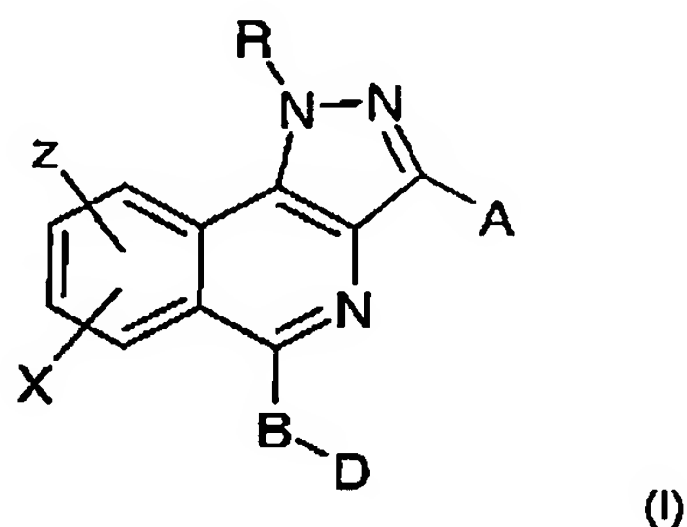
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rejection reactions on the part of the body against a transplanted organ or  
rejection reactions on the part of the transplanted organ against the body.

11. (currently amended) A pharmaceutical composition comprising a compound  
of the formula (I)



or a stereoisomeric form or a pharmaceutically acceptable salt of the  
compound of the formula I, wherein

A is  $-(C_1-C_6)$ -alkyl, in which alkyl is straight-chain or branched  
and is optionally substituted, once or more, independently of each  
other, by

$-O-R^1$  or

$-C(O)-OR^1$ ,

$-C(O)-NR^1R^1$ ,

$-C(O)-NR^1-SO_2R^1$ ,

$-NR^1R^1$ ,

$-CN$ , in which  $R^1$  is

hydrogen,

$-(C_1-C_6)$ -alkyl,

$-(C_6-C_{14})$ aryl or

fluoroalkyl of the formula  $-C_nH_xF_y$  or fluoroalkoxy of

the formula  $-OC_nH_xF_y$ , wherein  $n$  is an integer from 1

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to 6, x is an integer from 0 to 12, y is an integer from 1 to 13 and sum of x and y is  $2n + 1$ ,

-O-R<sup>1</sup>,

-SR<sup>1</sup>,

-S(O)-R<sup>1</sup>

-S(O)<sub>2</sub>-R<sup>1</sup>

-C(O)-OR<sup>1</sup>,

fluoroalkyl of the formula -C<sub>n</sub>H<sub>x</sub>F<sub>y</sub> or fluoroalkoxy of the formula -OC<sub>n</sub>H<sub>x</sub>F<sub>y</sub>, wherein n is an integer from 1 to 6, x is an integer from 0 to 12, y is an integer from 1 to 13 and sum of x and y is  $2n + 1$ ,

-C(O)-NR<sup>1</sup>R<sup>1</sup>,

-C(O)-NR<sup>1</sup>-SO<sub>2</sub>R<sup>1</sup>,

-NR<sup>1</sup>R<sup>1</sup>,

-CN,

monocyclic or polycyclic heteroaryl having from 5 to 14 ring members, which contains 1 to 5 heteroatoms as ring members, wherein heteroatoms are selected from N, O and S, and in which heteroaryl is unsubstituted or optionally substituted once or more, independently of each other, by R<sup>2</sup>, in which R<sup>2</sup> is

-(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

-OH,

-O-(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

halogen,

-N(R<sup>3</sup>)-R<sup>4</sup>, in which R<sup>3</sup> and R<sup>4</sup> are, independently of each other, hydrogen atom or -(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

fluoroalkyl of the formula -C<sub>n</sub>H<sub>x</sub>F<sub>y</sub> or fluoroalkoxy of the formula -OC<sub>n</sub>H<sub>x</sub>F<sub>y</sub>, wherein n is an integer from 1 to 4, x is an

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integer from 0 to 8, y is an integer from 1 to 9 and sum of x and y is  $2n + 1$ ,

-CN,

-SR<sup>1</sup>,

-S(O)-R<sup>1</sup>,

-S(O)<sub>2</sub>-R<sup>1</sup> or

-C(O)-NR<sup>1</sup>R<sup>1</sup>,

-(C<sub>3</sub>-C<sub>8</sub>)-cycloalkyl, in which cycloalkyl is unsubstituted or substituted, once or more, independently of each other, by R<sup>2</sup>, and R<sup>2</sup> is defined as above,

monocyclic or bicyclic heterocycle having from 5 to 12 ring members, which contains 1 to 5 heteroatoms as ring members, wherein heteroatoms are selected from N, O and S, and in which heterocycle is unsubstituted or substituted, once or more, independently of each other, by R<sup>2</sup>, and R<sup>2</sup> is defined as above,

B is a covalent bond,

-C=CR<sup>1</sup>-,

—C≡C—

-O(CH<sub>2</sub>)<sub>a</sub>-, in which a is an integer from 1 to 4,

O, S, NR<sup>2</sup>, -C(O)-, -NR<sup>2</sup>-C(O)-, -C(O)-NR<sup>2</sup>-, -NR<sup>2</sup>-SO<sub>2</sub>-, -SO<sub>2</sub>-NR<sup>2</sup>-, -NR<sup>2</sup>-C(O)-NR<sup>2</sup>-, and R<sup>2</sup> is defined as above, or

-(C<sub>1</sub>-C<sub>4</sub>)-alkylene, in which alkylene is straight-chain or branched and is optionally substituted, once or more, independently of each other, by R<sup>1</sup>, and R<sup>1</sup> is defined as above,

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D is  $-(C_1-C_6)$ -alkyl, in which alkyl is straight-chain or branched and is optionally substituted, once or more, independently of each other, by  $R^1$ , and  $R^1$  is defined as above,

monocyclic or polycyclic heteroaryl having from 5 to 14 ring members, which contains 1 to 5 heteroatoms as ring members, wherein heteroatoms are selected from N, O and S, and in which heteroaryl is unsubstituted or is substituted once or more, independently of each other, by  $R^2$ , and  $R^2$  is defined as above,

monocyclic or bicyclic heterocycle having from 5 to 12 ring members, which contains 1 to 5 heteroatoms as ring members, wherein heteroatoms are selected from N, O and S, and in which heterocycle is unsubstituted or substituted, once or more, independently of each other, by  $R^2$ , and  $R^2$  is defined as above,

$-(C_6-C_{14})$ -aryl, in which aryl is unsubstituted or substituted, once or more, independently of each other, by  $R^2$ , and  $R^2$  is defined as above, or

$-(C_3-C_6)$ -cycloalkyl, in which cycloalkyl is unsubstituted or substituted, once or more, independently of each other, by  $R^2$ , and  $R^2$  is defined as above, or

B-D is hydrogen,  
halogen,

fluoroalkyl of the formula  $-C_nH_xF_y$  or fluoroalkoxy of the formula  $-OC_nH_xF_y$ , wherein n is an integer from 1 to 4, x is an integer from 0 to 8, y is an integer from 1 to 9 and sum of x and y is  $2n + 1$ ,

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$-(CH_2)_a-Y-R^3$ , in which  $a$  is an integer from 1 to 4,  $Y$  is O, S,  $NR^2$ , and  $R^3$  is

$-(C_1-C_6)$ -alkyl,

$-(C_6-C_{14})$ -aryl,

$-(C_3-C_6)$ -cycloalkyl, and

$R$  is hydrogen,  
 $-(C_1-C_6)$ -alkyl, or  
 $-(C_6-C_{14})$ -aryl- $-(C_1-C_6)$ -alkyl, in which aryl is unsubstituted or substituted, once or more, independently of each other, by  $R^2$ , and  $R^2$  is defined as above, and

$X$  and  $Z$  are identical or different and are, independently of each other selected from:

hydrogen atom,

$-(C_1-C_4)$ -alkyl,

$-OH$ ,

$-O-(C_1-C_4)$ -alkyl),

halogen,

fluoroalkyl of the formula  $-C_nH_xF_y$  or fluoroalkoxy of the formula  $-OC_nH_xF_y$ , wherein  $n$  is an integer from 1 to 6,  $x$  is an integer from 0 to 12,  $y$  is an integer from 1 to 13 and sum of  $x$  and  $y$  is  $2n + 1$ ,

$-C(O)-OR^1$ ,

$-C(O)-NR^1R^1$ ,

$-C(O)-NR^1-SO_2R^1$ ,

$-NR^1R^1$ ,

$-NR^1-C(O)-NR^1R^1$ ,

$-NR^1-C(O)-R^1$ ,

$-NR^1-C(O)-OR^1$ ,

$-O-C(O)-NR^1R^1$ ,



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-CN,

-SR<sup>1</sup>,

-S(O)-R<sup>1</sup>,

-S(O)<sub>2</sub>-R<sup>1</sup>,

-S(O)<sub>2</sub>-NR<sup>1</sup>R<sup>1</sup>,

-NR<sup>1</sup>-SO<sub>2</sub>-R<sup>1</sup>, in which R<sup>1</sup> is as defined above,

monocyclic or bicyclic heterocycle having from 5 to 12 ring members, which contains 1 to 5 heteroatoms as ring members, wherein heteroatoms are selected from N, O and S, and in which heterocycle is unsubstituted or substituted, once or more, independently of each other, by R<sup>2</sup>, and R<sup>2</sup> is defined as above, or

-(C<sub>3</sub>-C<sub>6</sub>)-cycloalkyl, in which cycloalkyl is unsubstituted or substituted, once or more, independently of each other, by R<sup>2</sup>, and R<sup>2</sup> is defined as above,

with the proviso that when A is -(C<sub>1</sub>-C<sub>6</sub>)-alkyl, in which alkyl is straight-chain or branched and is optionally substituted, once or more, independently of each other, by

-O-R<sup>1</sup> or

-C(O)-OR<sup>1</sup>, in which R<sup>1</sup> is

hydrogen,

-(C<sub>1</sub>-C<sub>6</sub>)-alkyl,

-O-R<sup>1</sup>, in which R<sup>1</sup> is as defined in this proviso,

-C(O)-OR<sup>1</sup>, in which R<sup>1</sup> is as defined in this proviso, or

monocyclic or polycyclic heteroaryl having from 5 to 14 ring members, which contains 1 to 5 heteroatoms as ring members, wherein heteroatoms are selected from N, O and S, and in which heteroaryl is unsubstituted or optionally substituted once or more, independently of each other, by R<sup>2</sup>, in which R<sup>2</sup> is

hydrogen,

-(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

-OH,

-O-(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

halogen, or

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-N(R<sup>3</sup>)-R<sup>4</sup>, in which R<sup>3</sup> and R<sup>4</sup> are, independently of each other, hydrogen atom or -(C<sub>1</sub>-C<sub>4</sub>)-alkyl, then at least one of the following applies:

B is not a covalent bond or -(C<sub>1</sub>-C<sub>4</sub>)-alkylene;

~~D is not heteroaryl, heterocycle, (C<sub>6</sub>-C<sub>14</sub>)-aryl, (C<sub>3</sub>-C<sub>6</sub>)-cycloalkyl, or~~

~~X and Z are not (C<sub>1</sub>-C<sub>4</sub>)-alkyl, OH, O-(C<sub>1</sub>-C<sub>4</sub>)-alkyl, or halogen.~~

12. (original) The composition as claimed in claim 11, wherein

A is -(C<sub>1</sub>-C<sub>3</sub>)-alkyl, in which alkyl is straight-chain or branched and is optionally substituted, once or more, independently of each other, by

-O-R<sup>1</sup>, or

-C(O)-OR<sup>1</sup>, in which R<sup>1</sup> is

hydrogen,

-(C<sub>1</sub>-C<sub>3</sub>)-alkyl, or

-CF<sub>3</sub>

fluoroalkyl of the formula -C<sub>n</sub>H<sub>x</sub>F<sub>y</sub> or fluoroalkoxy of the formula -OC<sub>n</sub>H<sub>x</sub>F<sub>y</sub>, wherein n is an integer from 1 to 3, x is an integer from 0 to 6, y is an integer from 1 to 7 and sum of x and y is 2n + 1,

B is a covalent bond or O,

D is phenyl or naphthyl, in which phenyl or naphthyl is unsubstituted or substituted, once or more, independently of each other, by R<sup>2</sup>, in which R<sup>2</sup> is

fluorine, chlorine or bromine,

-OH,

-CF<sub>3</sub>,

-SR<sup>1</sup>, in which R<sup>1</sup> is defined as above,

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-(C<sub>1</sub>-C<sub>4</sub>)-alkyl  
-O-(C<sub>1</sub>-C<sub>2</sub>)-alkyl or  
-N(R<sup>3</sup>)-R<sup>4</sup>, in which R<sup>3</sup> and R<sup>4</sup> are, independently of each other, hydrogen atom or -(C<sub>1</sub>-C<sub>3</sub>)-alkyl,

heteroaryl selected from the group consisting of pyridyl, furanyl, pyrrolyl, isoxazolyl, benzofuranyl, benzothiophenyl, quinolinyl, isoquinolinyl, quinoxalinyl and thiophenyl, in which heteroaryl is unsubstituted or substituted, once or more, independently of each other, by R<sup>2</sup>, in which R<sup>2</sup> is defined as above or

-(C<sub>4</sub>-C<sub>6</sub>)-cycloalkyl, in which cycloalkyl is unsubstituted or substituted, once or more, independently of each other, by R<sup>2</sup>, and R<sup>2</sup> is defined as above, or

B-D is ((CH<sub>2</sub>)<sub>a</sub>-Y-R<sup>3</sup>), in which a is an integer from 1 to 2, Y is O and R<sup>3</sup> is -(C<sub>1</sub>-C<sub>3</sub>)-alkyl, and

R is hydrogen,

-(C<sub>1</sub>-C<sub>3</sub>)-alkyl, or  
-phenyl-(C<sub>1</sub>-C<sub>3</sub>)-alkyl, and

X and Z are identical or different and are, independently of each other, hydrogen, -C(O)-O(C<sub>1</sub>-C<sub>3</sub>)alkyl, -OCH<sub>3</sub>, -N(CH<sub>3</sub>)<sub>2</sub> or halogen.

13. (original) The composition as claimed in claim 11, wherein the compound of the formula I is selected from the group consisting of:

5-pyridin-2-yl-3-trifluoromethyl-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(2-trifluoromethyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(3-trifluoromethyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(4-trifluoromethyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,

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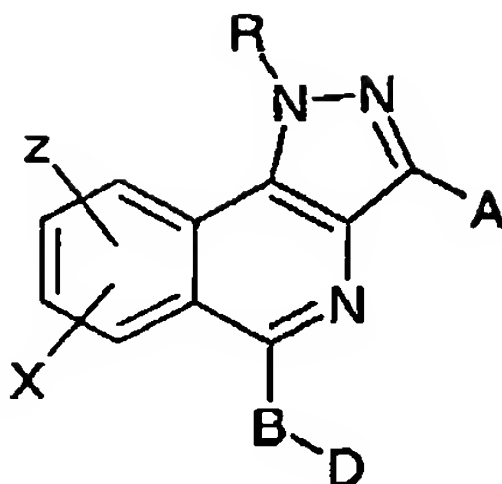
1,3-dimethyl-5-(3-trifluoromethylphenyl)-1H-pyrazolo[4,3-c]-isoquinoline,  
5-phenyl-3-trifluoromethyl-1H-pyrazolo[4,3-c]isoquinoline,  
1,3-dimethyl-5-(3-trifluoromethylphenyl)-1H-pyrazolo[4,3-c]-isoquinoline,  
1,3-dimethyl-5-(2,6-difluorophenyl)-1H-pyrazolo[4,3-c]-isoquinoline,  
1-benzyl-5-cyclohexyl-3-methyl-1H-pyrazolo[4,3-c]-isoquinoline,  
1-benzyl-5-naphthyl-3-methyl-1H-pyrazolo[4,3-c]-isoquinoline,  
5-methoxymethyl-3-methyl-1H-pyrazolo[4,3-c]-isoquinoline,  
7-methoxycarbonyl-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]-isoquinoline,  
7-methoxycarbonyl-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]-  
isoquinoline,  
7-dimethylamino-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
7-dimethylamino-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]-isoquinoline,  
6-dimethylamino-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
6-dimethylamino-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]-isoquinoline,  
8-dimethylamino-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
8-dimethylamino-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]-isoquinoline,  
1,3-dimethyl-5-(3-methyl-thiophen-2-yl)-1H-pyrazolo[4,3-c]-isoquinoline,  
3-methyl-5-phenyl-9-trifluoromethyl-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-pyridin-2-yl-9-trifluoromethyl-1H-pyrazolo[4,3-c]-isoquinoline,  
and  
3-methyl-5-(2,3,4,5,6-pentafluoro-phenyl)-1H-pyrazolo[4,3-c]-isoquinoline.

14. (currently amended) A method of treating a disease condition associated with inflammation comprising administering to a patient suffering from said disease condition a therapeutically effective amount of a compound of formula (I):

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(I)

or a stereoisomeric form or a pharmaceutically acceptable salt of the compound of the formula I, wherein

A is  $-(C_1-C_6)$ -alkyl, in which alkyl is straight-chain or branched and is optionally substituted, once or more, independently of each other, by

$-O-R^1$  or  
 $-C(O)-OR^1$ ,  
 $-C(O)-NR^1R^1$ ,  
 $-C(O)-NR^1-SO_2R^1$ ,  
 $-NR^1R^1$ ,  
 $-CN$ , in which  $R^1$  is

hydrogen,

$-(C_1-C_6)$ -alkyl,

$-(C_6-C_{14})$ aryl or

fluoroalkyl of the formula  $-C_nH_xF_y$  or fluoroalkoxy of the formula  $-OC_nH_xF_y$ , wherein  $n$  is an integer from 1 to 6,  $x$  is an integer from 0 to 12,  $y$  is an integer from 1 to 13 and sum of  $x$  and  $y$  is  $2n + 1$ ,

$-O-R^1$ ,

$-SR^1$ ,

$-S(O)-R^1$

$-S(O)_2-R^1$

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-C(O)-OR<sup>1</sup>,

fluoroalkyl of the formula -C<sub>n</sub>H<sub>x</sub>F<sub>y</sub> or fluoroalkoxy of the formula -OC<sub>n</sub>H<sub>x</sub>F<sub>y</sub>, wherein n is an integer from 1 to 6, x is an integer from 0 to 12, y is an integer from 1 to 13 and sum of x and y is 2n + 1,

-C(O)-NR<sup>1</sup>R<sup>1</sup>,

-C(O)-NR<sup>1</sup>-SO<sub>2</sub>R<sup>1</sup>,

-NR<sup>1</sup>R<sup>1</sup>,

-CN,

monocyclic or polycyclic heteroaryl having from 5 to 14 ring members, which contains 1 to 5 heteroatoms as ring members, wherein heteroatoms are selected from N, O and S, and in which heteroaryl is unsubstituted or optionally substituted once or more, independently of each other, by

R<sup>2</sup>, in which R<sup>2</sup> is

-(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

-OH,

-O-(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

halogen,

-N(R<sup>3</sup>)-R<sup>4</sup>, in which R<sup>3</sup> and R<sup>4</sup> are, independently of each other, hydrogen atom or -(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

fluoroalkyl of the formula -C<sub>n</sub>H<sub>x</sub>F<sub>y</sub> or fluoroalkoxy of the formula -OC<sub>n</sub>H<sub>x</sub>F<sub>y</sub>, wherein n is an integer from 1 to 4, x is an integer from 0 to 8, y is an integer from 1 to 9 and sum of x and y is 2n + 1,

-CN,

-SR<sup>1</sup>,

-S(O)-R<sup>1</sup>,

-S(O)<sub>2</sub>-R<sup>1</sup> or

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$\text{-C(O)-NR}^1\text{R}^1$ ,

$\text{-(C}_3\text{-C}_6\text{)-cycloalkyl}$ , in which cycloalkyl is unsubstituted or substituted, once or more, independently of each other, by  $\text{R}^2$ , and  $\text{R}^2$  is defined as above,

monocyclic or bicyclic heterocycle having from 5 to 12 ring members, which contains 1 to 5 heteroatoms as ring members, wherein heteroatoms are selected from N, O and S, and in which heterocycle is unsubstituted or substituted, once or more, independently of each other, by  $\text{R}^2$ , and  $\text{R}^2$  is defined as above,

B is a covalent bond,

$\text{-C=CR}^1\text{-}$ ,

$\text{—C}\equiv\text{C—}$  ,

$\text{-O(CH}_2\text{)}_a\text{-}$ , in which a is an integer from 1 to 4,

O, S,  $\text{NR}^2$ ,  $\text{-C(O)-}$ ,  $\text{-NR}^2\text{-C(O)-}$ ,  $\text{-C(O)-NR}^2\text{-}$ ,  $\text{-NR}^2\text{-SO}_2\text{-}$ ,  $\text{-SO}_2\text{-NR}^2\text{-}$ ,  $\text{-NR}^2\text{-C(O)-NR}^2\text{-}$ , and  $\text{R}^2$  is defined as above, or

$\text{-(C}_1\text{-C}_4\text{)-alkylene}$ , in which alkylene is straight-chain or branched and is optionally substituted, once or more, independently of each other, by  $\text{R}^1$ , and  $\text{R}^1$  is defined as above,

D is  $\text{-(C}_1\text{-C}_8\text{)-alkyl}$ , in which alkyl is straight-chain or branched and is optionally substituted, once or more, independently of each other, by  $\text{R}^1$ , and  $\text{R}^1$  is defined as above,

monocyclic or polycyclic heteroaryl having from 5 to 14 ring members, which contains 1 to 5 heteroatoms as ring members, wherein heteroatoms are selected from N, O and S, and in which



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heteroaryl is unsubstituted or is substituted once or more,  
independently of each other, by  $R^2$ , and  $R^2$  is defined as above,

monocyclic or bicyclic heterocycle having from 5 to 12 ring  
members, which contains 1 to 5 heteroatoms as ring members,  
wherein heteroatoms are selected from N, O and S, and in which  
heterocycle is unsubstituted or substituted, once or more,  
independently of each other, by  $R^2$ , and  $R^2$  is defined as above,

-( $C_6$ - $C_{14}$ )-aryl, in which aryl is unsubstituted or substituted,  
once or more, independently of each other, by  $R^2$ , and  $R^2$  is defined  
as above, or

-( $C_3$ - $C_8$ )-cycloalkyl, in which cycloalkyl is unsubstituted or  
substituted, once or more, independently of each other, by  $R^2$ , and  $R^2$   
is defined as above, or

B-D is hydrogen,  
halogen,

fluoroalkyl of the formula  $-C_nH_xF_y$  or fluoroalkoxy of the  
formula  $-OC_nH_xF_y$ , wherein n is an integer from 1 to 4, x is an integer  
from 0 to 8, y is an integer from 1 to 9 and sum of x and y is  $2n + 1$ ,

$-(CH_2)_a-Y-R^3$ , in which a is an integer from 1 to 4, Y is O, S,  $NR^2$ , and  
 $R^3$  is

-( $C_1$ - $C_6$ )-alkyl,  
-( $C_6$ - $C_{14}$ )-aryl,  
-( $C_3$ - $C_8$ )-cycloalkyl, and

R is hydrogen,

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-(C<sub>1</sub>-C<sub>6</sub>)-alkyl, or  
-(C<sub>6</sub>-C<sub>14</sub>)-aryl-(C<sub>1</sub>-C<sub>6</sub>)-alkyl, in which aryl is unsubstituted or substituted, once or more, independently of each other, by R<sup>2</sup>, and R<sup>2</sup> is defined as above, and

X and Z are identical or different and are, independently of each other selected from:

hydrogen atom,  
-(C<sub>1</sub>-C<sub>4</sub>)-alkyl,  
-OH,  
-O-(C<sub>1</sub>-C<sub>4</sub>-alkyl),  
halogen,  
fluoroalkyl of the formula -C<sub>n</sub>H<sub>x</sub>F<sub>y</sub> or fluoroalkoxy of the formula -OC<sub>n</sub>H<sub>x</sub>F<sub>y</sub>, wherein n is an integer from 1 to 6, x is an integer from 0 to 12, y is an integer from 1 to 13 and sum of x and y is 2n + 1,  
-C(O)-OR<sup>1</sup>,  
-C(O)-NR<sup>1</sup>R<sup>1</sup>,  
-C(O)-NR<sup>1</sup>-SO<sub>2</sub>R<sup>1</sup>,  
-NR<sup>1</sup>R<sup>1</sup>,  
-NR<sup>1</sup>-C(O)-NR<sup>1</sup>R<sup>1</sup>,  
-NR<sup>1</sup>-C(O)-R<sup>1</sup>,  
-NR<sup>1</sup>-C(O)-OR<sup>1</sup>,  
-O-C(O)-NR<sup>1</sup>R<sup>1</sup>,  
-CN,  
-SR<sup>1</sup>,  
-S(O)-R<sup>1</sup>,  
-S(O)<sub>2</sub>-R<sup>1</sup>,  
-S(O)<sub>2</sub>-NR<sup>1</sup>R<sup>1</sup>,  
-NR<sup>1</sup>-SO<sub>2</sub>-R<sup>1</sup>, in which R<sup>1</sup> is as defined above,

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heterocycle having from 5 to 12 ring members, in which heterocycle is unsubstituted or substituted, once or more, independently of each other, by  $R^2$ , and  $R^2$  is defined as above, or

-(C<sub>3</sub>-C<sub>6</sub>)-cycloalkyl, in which cycloalkyl is unsubstituted or substituted, once or more, independently of each other, by  $R^2$ , and  $R^2$  is defined as above,

with the proviso that when A is -(C<sub>1</sub>-C<sub>6</sub>)-alkyl, in which alkyl is straight-chain or branched and is optionally substituted, once or more, independently of each other, by

-O-R<sup>1</sup> or  
-C(O)-OR<sup>1</sup>, in which R<sup>1</sup> is  
hydrogen,

-(C<sub>1</sub>-C<sub>6</sub>)-alkyl,

-O-R<sup>1</sup>, in which R<sup>1</sup> is as defined in this proviso,

-C(O)-OR<sup>1</sup>, in which R<sup>1</sup> is as defined in this proviso, or

monocyclic or polycyclic heteroaryl having from 5 to 14 ring members, which contains 1 to 5 heteroatoms as ring members, wherein heteroatoms are selected from N, O and S, and in which heteroaryl is unsubstituted or optionally substituted once or more, independently of each other, by  $R^2$ , in which  $R^2$  is

hydrogen,

-(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

-OH,

-O-(C<sub>1</sub>-C<sub>4</sub>)-alkyl,

halogen, or

-N(R<sup>3</sup>)-R<sup>4</sup>, in which R<sup>3</sup> and R<sup>4</sup> are, independently of each other, hydrogen atom or -(C<sub>1</sub>-C<sub>4</sub>)-alkyl, then

B is not a covalent bond or -(C<sub>1</sub>-C<sub>4</sub>)-alkylene.

15. (original) The method as claimed in claim 14 wherein

A is -(C<sub>1</sub>-C<sub>3</sub>)-alkyl, in which alkyl is straight-chain or branched and is optionally substituted, once or more, independently of each other, by

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-O-R<sup>1</sup>, or  
-C(O)-OR<sup>1</sup>, in which R<sup>1</sup> is  
hydrogen,  
-(C<sub>1</sub>-C<sub>3</sub>)-alkyl, or  
-CF<sub>3</sub>

fluoroalkyl of the formula -C<sub>n</sub>H<sub>x</sub>F<sub>y</sub> or fluoroalkoxy of the  
formula -OC<sub>n</sub>H<sub>x</sub>F<sub>y</sub>, wherein n is an integer from 1 to 3, x is an  
integer from 0 to 6, y is an integer from 1 to 7 and sum of x and y is  
2n + 1,

B is a covalent bond or O,

D is phenyl or naphthyl, in which phenyl or naphthyl is  
unsubstituted or substituted, once or more, independently of each  
other, by R<sup>2</sup>, in which R<sup>2</sup> is

fluorine, chlorine or bromine,  
-OH,  
-CF<sub>3</sub>,  
-SR<sup>1</sup>, in which R<sup>1</sup> is defined as above,  
-(C<sub>1</sub>-C<sub>4</sub>)-alkyl  
-O-(C<sub>1</sub>-C<sub>2</sub>)-alkyl or  
-N(R<sup>3</sup>)-R<sup>4</sup>, in which R<sup>3</sup> and R<sup>4</sup> are, independently of  
each other, hydrogen atom or -(C<sub>1</sub>-C<sub>3</sub>)-alkyl,

heteroaryl selected from the group consisting of  
pyridyl, furanyl, pyrrolyl, isoxazolyl, benzofuranyl, benzothiophenyl,  
quinolinyl, isoquinolinyl, quinoxalinyl and thiophenyl, in which  
heteroaryl is unsubstituted or substituted, once or more,  
independently of each other, by R<sup>2</sup>, in which R<sup>2</sup> is defined as above or

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-(C<sub>4</sub>-C<sub>8</sub>)-cycloalkyl, in which cycloalkyl is unsubstituted or substituted, once or more, independently of each other, by R<sup>2</sup>, and R<sup>2</sup> is defined as above, or

B-D is ((CH<sub>2</sub>)<sub>a</sub>-Y-R<sup>3</sup>, in which a is an integer from 1 to 2, Y is O and R<sup>3</sup> is -(C<sub>1</sub>-C<sub>3</sub>)-alkyl, and

R is hydrogen,

-(C<sub>1</sub>-C<sub>3</sub>)-alkyl, or

-phenyl-(C<sub>1</sub>-C<sub>3</sub>)-alkyl, and

X and Z are identical or different and are, independently of each other, hydrogen, -C(O)-O(C<sub>1</sub>-C<sub>3</sub>)alkyl, -OCH<sub>3</sub>, -N(CH<sub>3</sub>)<sub>2</sub> or halogen.

16. (original) The method as claimed in claim 14, wherein the compound of formula (I) is selected from the group consisting of:

5-pyridin-2-yl-3-trifluoromethyl-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(2-trifluoromethyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(3-trifluoromethyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(4-trifluoromethyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
1,3-dimethyl-5-(3-trifluoromethylphenyl)-1H-pyrazolo[4,3-c]-isoquinoline,  
5-phenyl-3-trifluoromethyl-1H-pyrazolo[4,3-c]isoquinoline,  
1,3-dimethyl-5-(3-trifluoromethylphenyl)-1H-pyrazolo[4,3-c]-isoquinoline,  
1,3-dimethyl-5-(2,6-difluorophenyl)-1H-pyrazolo[4,3-c]-isoquinoline,  
1-benzyl-5-cyclohexyl-3-methyl-1H-pyrazolo[4,3-c]-isoquinoline,  
1-benzyl-5-naphthyl-3-methyl-1H-pyrazolo[4,3-c]-isoquinoline,  
5-methoxymethyl-3-methyl-1H-pyrazolo[4,3-c]-isoquinoline,  
7-methoxycarbonyl-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]-isoquinoline,  
7-methoxycarbonyl-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]-isoquinoline,

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7-dimethylamino-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
7-dimethylamino-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
6-dimethylamino-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
6-dimethylamino-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
8-dimethylamino-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
8-dimethylamino-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
1,3-dimethyl-5-(3-methyl-thiophen-2-yl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-phenyl-9-trifluoromethyl-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-pyridin-2-yl-9-trifluoromethyl-1H-pyrazolo[4,3-c]isoquinoline,  
and  
3-methyl-5-(2,3,4,5,6-pentafluoro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline.

17. (original) The method as claimed in claim 14, wherein the compound of formula (I) is selected from the group consisting of:

5-benzo[b]thiophene-2-yl-3-methyl-1H-pyrazolo[4,3-c]isoquinoline,  
7-bromo-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
7-bromo-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
5-(3-chloro-benzo[b]thiophen-2-yl)-3-methyl-1H-pyrazolo[4,3-c]-  
isoquinoline,  
5-(2-chloro-pyridin-3-yl)-3-methyl-1H-pyrazolo[4,3-c]isoquinoline,  
5-(6-chloro-pyridin-3-yl)-3-methyl-1H-pyrazolo[4,3-c]isoquinoline,  
3-ethyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
3-ethyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
3-ethyl-5-pyridin-3-yl-1H-pyrazolo[4,3-c]isoquinoline,  
5-furan-2-yl-3-methyl-1H-pyrazolo[4,3-c]isoquinoline,  
7-methoxy-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(3-methyl-benzofuran-2-yl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
6-chloro-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
6-fluoro-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,

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8-chloro-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
8-fluoro-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
9-chloro-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
9-fluoro-3-methyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
3,9-dimethyl-5-phenyl-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(2-methyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(3-methyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(4-methyl-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(2-bromo-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(3-bromo-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(2-chloro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(4-chloro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(2,4-dichloro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(3,4-dichloro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(2,6-dichloro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(2-fluoro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(4-fluoro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(2,4-difluoro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(2,6-difluoro-phenyl)-1H-pyrazolo[4,3-c]isoquinoline,  
6-chloro-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
6-fluoro-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
8-chloro-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
8-fluoro-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
9-chloro-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
9-fluoro-3-methyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
3,9-dimethyl-5-pyridin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-(1-methyl-1H-pyrrol-2-yl)-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-quinolin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-quinolin-3-yl-1H-pyrazolo[4,3-c]isoquinoline,  
3-methyl-5-quinoxalin-2-yl-1H-pyrazolo[4,3-c]isoquinoline,



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3-methyl-5-thiophen-2-yl-1H-pyrazolo[4,3-c]isoquinoline, and  
3-methyl-5-(3-methyl-thiophen-2-yl)-1H-pyrazolo[4,3-c]isoquinoline.

18. (original) The method as claimed in claim 14, wherein the disease condition is selected from the group consisting of multiple sclerosis, atherosclerosis, inflammatory bowel disease, Alzheimer's disease, stroke and diabetes.
19. (original) The method as claimed in claim 18, wherein the disease condition is multiple sclerosis.
20. (original) The method as claimed in claim 18, wherein the disease condition is atherosclerosis.
21. (original) The method as claimed in claim 18, wherein the disease condition is inflammatory bowel disease.
22. (original) The method as claimed in claim 18, wherein the disease condition is Alzheimer's disease.
23. (original) The method as claimed in claim 18, wherein the disease condition is stroke.
24. (original) The method as claimed in claim 18, wherein the disease condition is diabetes.